BIRCH, STEWART, KOLASCH & BIRC 1, LLP P.O. Box 747 • Falls Church, Virginia 22040-0747

PLEASE NOTE: YOU MUST COMPLETE THE FOLLOWING

Telephone: (703) 205-8000 • Facsimile: (703) 205-8050

COMBINED DECLARATION AND POWER OF ATTORNEY FOR PATENT AND DESIGN APPLICATIONS

As a below named inventor, I hereby declare that: my residence, post office address and citizenship are as stated next to my name; that I verily believe that I am the original, first and sole inventor (if only one inventor is named below) or an original, first and joint inventor (if plural inventors are named below) of the subject matter which is claimed and for which a patent is sought on the invention entitled:

nsert Title:	PRODRUGS									
	the specification of which is attached hereto. If not attached hereto,									
ill in Appropriate nformation -	the specification was filed on February 12, 1999									
or Use Without	The state of the s									
pecification	and amended on					applicable)I	and/or			
ttached:	the specification was filed on						and was			
	United States Application Number 09/249.317 and amended on the specification was filed on International Application Number amended under PCT Article 19 on					(if ap	olicable)			
	I hereby state that I have reviewed and understand the contents of the above-identified specification, including the claims, as									
	1. I have a war and an our transformed to about									
	I acknowledge the duty to disclose information which is material to patentiability as defined in Title 37, Code of redefan									
	Regulations, §1.56. I do not know and do not believe the same was ever known or used in the United States of America before my or our invention.									
	thereof, or patented or described in any printed publication in any country before my or our invention thereof or more than one year prior to this application, that the same was not in public use or on sale in the United States of America more than one year prior to this application, that the same was not in public use or on sale in the United States of America more than one year									
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į.i.	year prior to this application, that the invention has not been patented or made the subject of an inventor's certificate issued before the date of this application, that the invention has not been patented or made the subject of an inventor's certificate issued before the date of this application, and that no application and that no application are the date of this application and that no application are the date of this application and that no application are the date of this application and that no application are the date of this application and that no application are the date of this application and that no application are the date of this application and that no application are the date of this application and the date of this application are the date of this application and the date of this application are the date of the date of this application are the date of this application are the date of the date of this application are the date of									
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	patent or inventor's certificate on this invention has been filed in any country foreign to the Office of America prior to this									
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	or inventor's certific	ate listed below a	nd have also identified beig tion on which priority is cla	w any foreign application imed:	101 patent of nev	eritor 5 cere	meate im ing			
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	98/7267		rica	August 13, 1998 (Month/Day/Year File	<u>d)</u>	⊠ Yes	□ No			
	(Number)	(Country		` '						
	I hereby claim the benefit under Title 35, United States Code, §119(e) of any United States provisional applications(s) listed below.									
	,									
Insert Provisional	/ 1: · · · · · · · · · · · · · · · · · ·			(Filing Date)						
Application(s):	(Application Numb	er)		(1 mile Date)						
(if any)										
	(Application Number) (Filing Date)									
-	All Foreign Applications, if any, for any Patent or Inventor's Certificate Filed More than 12 Months (6 Months for Designs) Prior to									
	the Filing Date of This Application:									
	D (* 457)					v/Year)				
	Country		Application Number	Date of 1	iling (Month/Da	-,,,				
Insert Requested	•									
Information:										
(if appropriate)										
	I hereby claim the benefit under Title 35, United States Code, \$120 of any United States and/or PCT application(s) listed below and									
	insofar as the subject matter of each of the claims of this application is not disclosed in the first paragraph of Title 35 United States Code. \$112.1 acknowledge the duty to disclose									
	application in the manner provided by the first paragraph of Title 35, Officer States Code, \$12,1 acts \$1.26 which became available information which is material to the patentability as defined in Title 37, Code of Federal Regulations, \$1.56 which became available between the filing date of the prior application and the national or PCT international filing date of this application.									
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In a and Dai II C	PCT/SE98/01467		August 14, 1998	pending			_,,.			
Insert Prior U.S. Application(s):	(Application Numb	per)	(Filing Date)	(Status -	patented, pendi	ng, abando	ned)			
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7	(Application Numb	oer)	(Filing Date)	(Status -	patented, pendi	ig, availuu	ica,			
Page 1 of 3										

I hereby appoint the t__owing attorneys to prosecute this application and, an international application based on this application and to transact all business in the Patent and Trademark Office connected therewith and in connection with the resulting patent based on instructions received from the entity who first sent the application papers to the attorneys identified below, unless the inventor(s) or assignee provides said attorneys with a written notice to the contrary:

Raymond C. Stewart	(Reg. No. 21,066)	Terrell C. Birch	(Reg. No. 19,382)
Joseph A. Kolasch	(Reg. No. 22,463)	James M. Slattery	(Reg. No. 28,380)
Bernard L. Sweeney	(Reg. No. 24,448)	Michael K. Mutter	(Reg. No. 29,680)
Charles Gorenstein	(Reg. No. 29,271)	Gerald M. Murphy, Jr.	(Reg. No. 28,977)
Leonard R. Svensson	(Reg. No. 30,330)	Terry L. Clark	(Reg. No. 32,644)
Andrew D. Meikle	(Reg. No. 32,868)	Marc S. Weiner	(Reg. No. 32,181)
Joe McKinney Muncy	(Reg. No. 32,334)	Donald J. Daley	(Reg. No. 34,313)
John W. Bailey	(Reg. No. 32,881)	John A. Castellano	(Reg. No. 35,094)

Send Correspondence to:

BIRCH, STEWART, KOLASCH & BIRCH, LLP

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PLEASE NOTE: YOU MUST COMPLETE THE FOLLOWING:

Full Name of First or Sole Inventor: Insert Name of Inventor Insert Date This Document is Signed

Insert Kesicence Insert Citizenship

ert Post Office

Full Name of Second

Full Name of Third

Full Name of Fourth

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I hereby declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code and that such willful false statements may jeopardize the validity of the application or any patent issued thereon.

DATE* INVENTOR'S SIGNATURE GIVEN NAME/FAMILY NAME -10-28 Xiao-Xiong ZHOU CITIZENSHII Residence (City, State & Country) Swedish Huddinge, Sweden POST OFFICE ADDRESS (Complete Street Address including City, State & Country) Kallkarsvagen 12, 141 41 Huddinge, Sweden INVENTOR'S SIGNATURE GIVEN NAME/FAMILY NAME 991020 Nils Gunnar JOHANSSON CITIZENSHIP Residence (City, State & Country) Swedish Enhorna, Sweden POST OFFICE ADDRESS (Complete Street Address including City, State & Country) Baverstigen 19, 150 23 Enhorna, Sweden DATE* INVENTOR'S SIGNATURE GIVEN NAME/FAMILY NAME *991020* Ment alex Horst WAHLING CITIZENSHIP Residence (City, State & Country) German Skarholmen, Sweden POST OFFICE ADDRESS (Complete Street Address including City, State & Country) Tempelriddarvagen 3, 127 61 Skarholmen, Sweden GIVEN NAME/FAMILY NAME INVENTOR'S SIGNATURE 991028 Christian SUND CITIZENSHIE Residence (City, State & Country) Finnish Varby, Sweden POST OFFICE ADDRESS (Complete Street Address including City, State & Country) Backgadsvagen 16, 2tr, 143 42 Varby, Sweden DATE* INVENTOR'S SIGNATURE GIVEN NAME/FAMILY NAME 991020 Hans WALLBERG CITIZENSHIP Residence (City, State & Country) Swedish Huddinge, Sweden POST OFFICE ADDRESS (Complete Street Address including City, State & Country) Mangardsvagen 10, SE-141 51 Huddinge, Sweden

Page 2 of 3 (Revised 11-98)

^{*}DATE OF SIGNATURE

ne of Sixth or, if any:	GIVEN NAME/FAMILY NAME	INVENTOR'S SIGNATURE		DATE*					
see above	Lourdes SALVADOR	Lousder Q. Johns	da	Ordoler 20, 1989					
	Residence (City, State & Country)	1	CITIZENSH	IP					
	Norrkoping, Sweden		Filipi	ino					
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Name of Seventh ventor, if any:	GIVEN NAME/FAMILY NAME	INVENTOR'S SIGNATURE		DATE*					
see above	Stefan LINDSTROM	Steam I		991101					
	Residence (City, State & Country)		CITIZENSH						
	Uppsala, Sweden	Swedish							
	POST OFFICE ADDRESS (Complete Street Address including City, State & Country)								
	Marmorsvagen 8B, SE-752 44 Uppsala, Sweden								
ı	GIVEN NAME/FAMILY NAME	INVENTOR'S SIGNATURE	I i	DATE*					
bove	Christer SAHLBERG	I Club Ja	4	October 20, 1999					
	Residence (City, State & Country)		CITIZENSH	¥ .					
	Hagersten, Sweden		Swedish						
	POST OFFICE ADDRESS (Complete Street	Address including City, State & Countr	y)						
	Malarhojdsvagen 5L, SE-129 40 Hagersten, Sweden								
	GIVEN NAME/FAMILY NAME	INVENTOR'S SIGNATURE ,		DATE*					
ve		Ch. A. Sat							
	Residence (City, State & Country)		CITIZENSH	IP					
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	POST OFFICE ADDRESS (Complete Street Address including City, State & Country)								

Page 3 of 3 (Revised 11-98)

*DATE OF SIGNATURE

PATENT Attorney Docket No. 1718-163P

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

APPLICANTS:

ZHOU, Xiao-Xiong et al.

SERIAL No.:

09/249,317

GROUP:

1623

FILED:

February 12, 1999

EXAMINER: J. Wilson

FOR:

PRODRUGS

DECLARATION SUBMITTED UNDER 37 C.F.R. § 1.132

Honorable Commissioner Of Patents and Trademarks Washington, D.C. 20231

Sir:

I, Dr. Xiao-Xiong Zhou Medivir AB, Lunastigen 7, S-141 44 Huddinge, Sweden, do hereby declare the following:

I have attached a copy of my curriculum vitae to this Declaration.

I am Director of Medicinal Chemistry at Medivir AB, a pharmaceutical research company employing around 90 research scientists, of which 53 are PhDs. I am familiar with the above referenced patent application, as well as the development, usages and properties of prodrug compounds.

I have read and understand the subject matter of the Office Action of January 31, 2001.

SERIAL No.: 09/249,317

The following comments are offered in support of the patentability of the instant invention.

The empty bracket symbol, "()_x", where x is 0, 1, 2, etc., that is used throughout the Specification and Claims of the instant application is a widely used convention for methylene homologues; i.e., for homologues varying by the presence of one or more contiguous –CH₂– units. In the case where x is zero, there is no methylene group present and the empty bracket then represents a bond. One skilled in the art of medicinal chemistry, particularly prodrugs, would easily understand the meaning of the empty bracket symbol.

In the appendix to this Declaration I have attached a number of examples taken from the Derwent patent abstracting service and that show the empty bracket symbol in use. Note that the authors of such abstracts regard the symbol as so clear that no further definition of it is present, or required, in the key.

In my opinion, no medicinal chemist would be in doubt as to the significance of the empty bracket symbol in the context of a patent Specification describing and exemplifying prodrugs of active compounds, where the prodrug moiety comprises (in part) unbranched alkyl chains (i.e. methylene homologues) of various lengths.

However, if it were desirable to avoid using shorthand conventions for the intended homologues, then I believe that it is crystal clear that the symbol "(CH₂)_x" should be put in the place of "()_x".

SERIAL No.: 09/249,317

The undersigned hereby declares that all statements made herein based upon knowledge are true, and that all statements made based upon information and belief are believed to be true; and further, that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code, and that such willful false statements may jeopardize the validity of the application or any patent issued thereon.

DATED:27 April 2001

Dr. Xiao-Xiong Zhou

Enclosures: As stated above